Dosimetric Comparison between Conventional 2D and 3D Conformal Radiotherapy in the Treatment of Intact Breast Cancer

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Abstract

Background: Radiotherapy (RT) techniques after Conservative Breast Surgery (CBS) vary. Three Dimension (3D) planning allows for better plan optimization compared to 2 Dimension (2D) plans and also allowing for creating Dose Volume Histograms (DVHs) for both Planning Target Volume (PTV) and Organs at Risk (OAR).

Patients and Methods: Twenty consecutive patients with CBS planned for whole breast and supraclavicular (SCV) RT at the National Cancer Institute (NCI), Egypt between January and June 2016 were included in this study. All patients were planned clinically in 2D fashion with no more than 2 cm of ipsilateral lung allowed in the tangential fields “Limited 2D” (Limit-2D) then Target and OAR volumes were drawn according to the Radiation Therapy Oncology Group (RTOG) guidelines and 3D plans and a central slice PTV-based 2D plan, “Modified 2D” (Mod-2D), were performed in the same Computerized Tomography (CT) slices for each patient. Mono-Iso-Centeric technique (MIT) was used in 3D plans. DVH parameters were used to compare the three plans. Results: In 3D plans, compared to Limit-2D, coverage improved for the intact breast (V95% = 95% versus (Vs) 69%, p = 0.036) and SCVPTV (V90% = 90% Vs 65%, p = 0.01). The breast and SCV V107%, V112% and Dmax were better with 3D plan however not statistical significant (NS). Junctional hot spots were 120% and 107% in the Limit-2D and 3D plans respectively (p = 0.04). The dose to the heart, mean (333 Vs 491 cGy), V10 (5% Vs 10%) and V20 (3% Vs 7%), Ipsilateral lung V20 (19% Vs 26%), and contra lateral breast D-max (205 Vs 462 cGy) were higher in 3D plans however NS, and the dose to the cord was the same. Comparison be-
tween 3D and Mod-2D showed better OAR sparing with 3D with mean heart dose (491 cGy Vs 782 cGy, p = 0.025) and Ipsilateral lung V20 (26% Vs 32%, p = 0.07% with statistically comparable target coverage. **Conclusion:** This study demonstrated that application of 3D plan using MIT improves coverage of breast and SCVPTVs with minimizing hot spot at the junctional area if compared with Limit-2D plans with comparable dose to OAR. When compared with Mod-2D plans, 3D plans not only had better target coverage but also better sparing of OAR, the latter was statistically significant.

**Keywords**
Mono-Iso-Centeric Technique, Breast Contouring, Breast Conserving Radiotherapy, Dose Volume Histogram (DVH)

**1. Introduction**

Breast cancer is the most common cause of cancer death among women worldwide. In Egypt, breast cancer is the most common cancer among women, representing 17.5% of total cancer cases (34.3% in Women and 0.5% in men) among the National Cancer Institute (NCI) series of 9808 patients during the year 2001, 2003 and 2004, Nadia Mokhtar et al., [1]. After CBS, RT of the breast is performed in most RT centers by using two simple tangential beams. This can be done by using either 2D or 3D treatment planning. In case of 2D treatment planning, the breast is treated by using two tangential parallel opposed fields and one anterior field to treat SCV lymph nodes if needed. The borders of the two tangential beams are determined clinically and the breast is manually contoured in the single axial slice where calculation is done, and SCV lymph nodes are calculated at depth 3 cm, Chika N. Madu et al., [2].

But in case of 3D treatment planning, full CT slices are used to treat both breast and SCV lymph nodes. The use of Multi-Leaf Collimator (MLC) and fields segments allows for better plan optimization. Usage of full CT slices allows for 3D evaluation of dose distribution, minimum, maximum dose and dose to OAR using DVH.

The use of 3D conformal radiotherapy and Intensity Modulated Radiation Therapy (IMRT) in breast cancer was associated with improved acute toxicity and cosmesis [3] [4], in addition to better target coverage DVH parameters [5], however; in case of IMRT, this was associated with higher dose to contra lateral OAR with the known risk of secondary malignancy [6].

With the higher patient-machine ratio and the limited availability of MLC-equipped treatment machines in Egypt and most developing countries, even with the use of hypo-fractionated breast RT schedules, every effort should be made to settle on a safe and simpler RT technique.

The aim of this study was to compare dosimetrically between 3D treatment planning using MIT and 2 methods of 2D treatment planning for patients with breast cancer that underwent conservative surgery with respect to PTV coverage
of the breast and the SCV lymph node, hot spot in the junctional area and dose to OAR.

2. Patients and Methods

2.1. Patients Selection

Twenty consecutive patients with CBS, in whom whole breast and SCV field irradiation were indicated and were treated at NCI Egypt between January and June 2016, were included in this study. Ten of them had right breast cancer and the other ten patients had left breast cancer.

2.2. RT Procedures

All patients were planned clinically using anatomical and bony landmarks. Radiopaque wire ring was placed around the breast to define the breast borders. Patient underwent CT scanning with 25 mm slice thickness. Target (breast and SCV lymph nodes) and OAR (i.e. heart in case of left sided breast, ipsilateral lung, contra-lateral breast and spinal cord) volumes were delineated according to RTOG guidelines [7]. Treatment planning was done on Precise Treatment Planning System.

2.3. Two Dimension Plans

2.3.1. Limited 2D (Limit-2D) Plan

Borders of the two tangential parallel opposed beams were defined clinically. The superior border covers as much breast tissue as possible and lies at the lower border of medial end of clavicle. The inferior border lies 2 cm below the inframammary fold, the medial border is usually in the midline, and the lateral border is in the mid-axillary line, Helen McNair et al., [8]. The posterior borders of the tangential (Tang) fields are aligned to each other and allowed to include not more than 2 cm of the ipsilateral lung (an institutional method that was frequently used to decrease lung toxicity). This plan was considered limited 2D plan.

2.3.2. Modified 2D (Mod-2D) Plan

Re-planning was done in which the 2 tang fields covered the central slice PTV properly provided no crossing of middle line and regardless of how much of the ipsilateral lung was included.

2D Plan Calculations Were Performed in the Central CT-Slice. Wedges were used to improve tissue dose homogeneity and all patients were treated with 6-MV photon beam except in one patient 15 MV photon beam was used.

2.3.3. SCV lymph Node Field

For SCV Lymph Node Field, Borders Were Defined Anatomically. The superior border has extended to the thyrocricoid membrane, inferior border has extended to the inferior aspect of the clavicular head matching with tangential field, medial border has extended to the midpoint, and lateral border has extended to the humeral head [2]. Calculation was done in the central CT-slice at depth 3 cm using single anterior field at SSD 100 cm with energy equal 6 MV and gantry an-
gle equal 10 degree to avoid the spinal cord.

2.4. Three Dimension Conformal Planning

In the 3D treatment planning Clinical Target Volume (CTV) and PTV of the breast and SCV lymph nodes were delineated according to RTOG guidelines [7]. MIT was applied to avoid the divergence between tangential beams of breast and the SCV field(s). In this technique the center of all beams was placed at the junction between the PTV of the breast and the PTV of SCV, Svensson GK et al., [9]. The suitable energy, 6 MV or 15 MV was used in these beams. The gantry angles were determined using the Beam’s Eye View (BEV). MLC was used to conform the prescribed dose around the PTVs with a margin of 0.7 cm to avoid the penumbra. Breast field segments were used instead of wedges for plan optimization. In PTV SCV, posterior fields were allowed. The planning was performed based on the 3D Algorithm using Precise-TPS.

2.5. Prescribed Dose and Plan Evaluation

A dose of 4005 cGy in 15 fractions over 3 weeks was prescribed. In all techniques DVHs were used to determine V95% (PTV receiving 95% of prescribed dose), V90%, V107%, V112%, maximum dose (D-max), and hot spots according to International Commission on Radiation Unit and Measurements (ICRU 50). [10], heart mean dose, V10 (volume of the heart receiving 10 Gy) and V20 in case of left sided breast, the volume of Ipsilateral lung receiving 20 Gy (V20), contra-lateral breast D-max and spinal cord D-max were derived and compared according to RTOG 1005 guidelines [11].

2.6. Statistical Analysis

SPSS version 22.0 software (Chicago, IL, USA) was used. The DVH parameters of the cumulative dose plans were compared with analysis of the mean values with the paired-samples t-test. All tests were two-tailed, and differences were considered statistically significant at $p \leq 0.05$.

3. Results

3.1. Limit-2D and 3D Plan Comparison

The Limit-2D plan, where post borders of the 2 tangential fields were set to include not more than 2 cm of the ipsilateral lung at the central slice, was compared to 3D plan. The latter used MIT.

The mean V95%, V107%, V112%, and D-max for the breast and mean V90%, V107%, V112%, and D-max for the SCV were better with 3D plans compared to Limit-2D plans and was statistically significant for V95% breast and V90% SCV (p 0.036 and 0.01 respectively) (Table 1). The Limit-2D plans had lower dose to OAR however without statistical significance (Table 1). Hot spot at the junction between the breast and SCV field in Limit-2D plans had a mean dose of 120% compared to 107% in 3D plans which was statistically significant (Table 1).
Table 1. The dosimetric comparison between limit-2D* and 3D** plans.

<table>
<thead>
<tr>
<th>Parameter</th>
<th>Limit-2D Plan</th>
<th>3D Plan</th>
<th>P Value</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Mean (Std. Deviation)</td>
<td>Mean (Std. Deviation)</td>
<td></td>
</tr>
<tr>
<td>Breast PTV***</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>V95%</td>
<td>69% (20%)</td>
<td>95% (3%)</td>
<td>0.036</td>
</tr>
<tr>
<td>V107%</td>
<td>8.2% (6%)</td>
<td>2.8% (2.5%)</td>
<td>0.12</td>
</tr>
<tr>
<td>V112%</td>
<td>0.5 % (0.5%)</td>
<td>0%</td>
<td>0.55</td>
</tr>
<tr>
<td>D-max</td>
<td>124 % (7%)</td>
<td>111% (1.7%)</td>
<td>0.08</td>
</tr>
<tr>
<td>V90%</td>
<td>65% (18%)</td>
<td>90% (4%)</td>
<td>0.01</td>
</tr>
<tr>
<td>Supraclavicular PTV</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>V90%</td>
<td>10% (7%)</td>
<td>4 (2%)</td>
<td>0.15</td>
</tr>
<tr>
<td>V112%</td>
<td>3% (2%)</td>
<td>1% (1%)</td>
<td>0.12</td>
</tr>
<tr>
<td>D-max</td>
<td>123% (8%)</td>
<td>113% (4%)</td>
<td>0.11</td>
</tr>
<tr>
<td>Junctional Hot Spot</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Mean</td>
<td>333cGy (190 cGy)</td>
<td>491cGy (110 cGy)</td>
<td>0.14</td>
</tr>
<tr>
<td>Heart in Left Side Breast (10 Patients)</td>
<td>V10 5% (4%)</td>
<td>10% (7%)</td>
<td>0.13</td>
</tr>
<tr>
<td>V20</td>
<td>3% (3%)</td>
<td>7% (5%)</td>
<td>0.14</td>
</tr>
<tr>
<td>Ipsilateral Lung V20</td>
<td>19% (8%)</td>
<td>26% (6%)</td>
<td>0.15</td>
</tr>
<tr>
<td>Contra-Lat Breast D-max</td>
<td>205 cGy (137 cGy)</td>
<td>462 cGy (144 cGy)</td>
<td>0.25</td>
</tr>
<tr>
<td>Spinal Cord</td>
<td>D-max</td>
<td>2457 cGy</td>
<td>2318 cGy</td>
</tr>
</tbody>
</table>

*Limit-2D where post border was set to include not more than 2 cm of ipsilateral lung at central slice. **3D using Mono-Iso-centric Technique (MIT). ***PTV (Planning Target Volume).

3.2. Comparison between Mod-2D Plan and 3D Plan

In the Mod-2D plan the 2 tang fields covered the central slice breast PTV properly regardless of how much Ipsilateral lung was included. This improved breast PTV V95% however on the expense of less sparing of OAR. The mean heart dose was statistically significantly lower in 3D plan compared to the Mod-2D plan. Also there was a trend towards significance in favor of 3D plan regarding breast D-max, heart V10 and Ipsilateral lung V20 (Table 2). SCV field and junctional hot spot were the same difference as with Limit-2D plan.

4. Discussion

For decades, patients with breast cancer received post operative RT using 2D technique. This technique didn’t allow the radiation oncologist and physicist to know dose to OAR or to the PTV above and below the central slice and dose variation across junction area between breast and SCV fields. This study aims at evaluating two 2D techniques and to compare it to 3D planning with the use of MIT.

4.1. Target Coverage

In case of the 3D treatment planning, breast V95% was 95% (±3%) compared to
Table 2. The comparison between Mod-2D* and 3D** plans.

<table>
<thead>
<tr>
<th>Parameter</th>
<th>Mod-2D Plan Mean (Std. Deviation)</th>
<th>3D Plan Mean (Std. Deviation)</th>
<th>P Value</th>
</tr>
</thead>
<tbody>
<tr>
<td>Breast PTV***</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>V95%</td>
<td>84% (10%)</td>
<td>95% (3%)</td>
<td>0.29</td>
</tr>
<tr>
<td>V107%</td>
<td>7.3% (4%)</td>
<td>2.8% (2.5%)</td>
<td>0.11</td>
</tr>
<tr>
<td>V112%</td>
<td>0.9% (0.9%)</td>
<td>0%</td>
<td>0.54</td>
</tr>
<tr>
<td>D-max</td>
<td>115% (3%)</td>
<td>111% (1.7%)</td>
<td>0.06</td>
</tr>
<tr>
<td>Heart in Left Side Breast (10 Patients)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Mean</td>
<td>782 cGy (100 cGy)</td>
<td>491 cGy (110 cGy)</td>
<td>0.025</td>
</tr>
<tr>
<td>V10</td>
<td>25% (7%)</td>
<td>10% (7%)</td>
<td>0.07</td>
</tr>
<tr>
<td>V20</td>
<td>8% (5%)</td>
<td>7% (5%)</td>
<td>0.2</td>
</tr>
<tr>
<td>Ipsilateral Lung</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>V20</td>
<td>32% (4%)</td>
<td>26% (6%)</td>
<td>0.07</td>
</tr>
<tr>
<td>Contra-Lat Breast</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>D-max</td>
<td>637 cGy (296)</td>
<td>462 cGy (144 cGy)</td>
<td>0.54</td>
</tr>
<tr>
<td>Spinal cord</td>
<td>D-max</td>
<td>979 cGy</td>
<td>1218 cGy</td>
</tr>
</tbody>
</table>

*The Mod-2D plan was modified so that the 2 tang fields covered the central slice breast PTV. **3D using Mono-Isocentric Technique (MIT). ***PTV (Planning Target Volume).

an average value of 69% (±20%) in Limit-2D plans. This better coverage was statistically significant (p = 0.036). Limiting post border of tangential fields to include not more than 2 cm of ipsilateral lung in these Limit-2D plans led to missing medial and lateral parts of the breast PTV and hence poor coverage. The 3D plans were also associated with better breast V107%, V112%, and D-max however NS.

The SCV V90% was 90% (±4%) with 3D compared with only 65% (±18%) in Limit-2D plan, again the difference was statistically significant (p = 0.01). The traditional use of 1 direct field for the SCV and calculation at depth 3 cm simply underestimates the depth of SCV and axillary apex lymph nodes specially in patients with high Body Mass Index (BMI) or big separation hence the poor coverage while in 3D posterior field was allowed with small weighting. With 3D plans SCV V107%, V112% and D-max were all lower than 2D plans however did not reach statistical significance.

The use of MIS in 3D plans allowed for marked elimination of junctional hot spots, 107% (±4%) compared to 120% (±9%) in 2D plans (p = 0.04).

In our study, the target coverage was better than that reported by Z. Falahatpouret et al., [12] (breast V95% was 74% with 2D and 81% with 3D plans) and inferior to Hans Paul van der Laan et al. [13] and Rudra et al. [5] whose results were similar (breast V95% was 95% with 2D and 99% with 3D plans). This inferior coverage with Limit-2D in our study can be explained by missing the medial and lateral Parts of the Breast (PTV) when the posterior border of the field was not allowed to include more than 2 cm of ipsilateral lung. When 2D plan was modified to cover breast PTV at central slice, Mod-2D, the coverage improved to 84% (±10%) but on the expense of more dose to OAR. SCV V90% in our study (65% in 2D and 90% in 3D) was slightly inferior to those reported by Rudra et
al., [5] (78% for 2D and 93.6% in 3D). Different patients’ average BMI or separation is a possible explanation.

In 2D plan the mean dose received in the junction area was (120% ± 9%) of the prescribed dose compared to (107% ± 4%) in 3D plans. This was comparable with Assaoui, F. et al. [14].

4.2. OAR Sparing

4.2.1. The Heart in Case of Left Sided Breast

In our study the average value of mean heart dose in case of left sided breast was 491 cGy in 3D plans compared to 782 cGy in Mod-2D plans and the difference was significant statistically ($p = 0.025$). Heart V10 and V20 were also better with 3D plan however NS. Although Limit-2D plans had statistically NS lower mean heart dose (333 cGy), this plan failed to cover the target volume properly.

This 3D mean heart dose is acceptable by the RTOG 1005 guidelines for left side breast and better than that reported by Rudra et al., (640 cGy) [5] and Hans Paul van der Laan et al., (550 cGy) [10]. Likewise the V10 and V20 in our study (10% and 7%) were better than reported by Rudra et al. (17.5% and 8%) This can be explained by the lower but still acceptable breast V95% in our study (95.3% compared to 99.2% in Rudra et al., study).

4.2.2. The Ipsilateral Lung

In our study Ipsilateral lung V20 was 26% in 3D plans compared to 32% in Mod-2D plans and there was trend towards statistical significance ($p = 0.07$). Although Limit-2D plans had statistically NS lower ipsilateral lung V20 (19%), this plan failed to cover the target volume properly.

This 3D Ipsilateral lung V20 is comparable with QUANTEC [15] guideline which recommended that V20 must deliver ($\leq 30\%$) and higher than accepted by the RTOG 1005 guidelines ($\geq 20\%$) however it is better than that reported by Rudra et al., (44%) [5]. This again can be explained by the lower but still acceptable breast V95% in our study (95.3% compared to 99.2% in Rudra et al., study).

Both Hans Paul van der Laan et al., [13], and Falahatpour et al., [12] were reporting the Ipsilateral lung V30 (3.5% and 9% respectively in 3D plans).

Further attempt to lower the dose to OAR was made in this study by lowering the breast PTV coverage to V90% $\geq 90\%$ instead of V95% $> 95$ (still accepted by RTOG 1005 guidelines), the Ipsilateral lung V20 dropped to 17% (compared to 26% with p value of 0.026) and mean heart dose dropped to 298 cGy (compared to 491 cGy with p value 0.09).

5. Limitations of the Study

As with similar type of studies, the clinical effect of the dosimetric findings cannot be evaluated. Those patients need to be followed up clinically for an appropriate time to assess for the difference in loco-regional control and late toxicity.

6. Conclusions

This study demonstrated that application of 3D plan using MIT improves cov-
verage of intact breast and SCV PTVs with elimination of hot spot at the junc-
tional area if compared with clinically-based Limit-2D plans however with non
statistically significant higher dose to OAR.

When comparing 3D plans with central slice PTV-based 2D (Mod-2D) plans,
3D plans not only had better target coverage but also better sparing of OAR, the
latter was statistically significant.

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